

**AMENDMENTS TO THE SPECIFICATION**

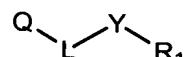
**Amend the specification by adding before the first line the sentence:**

This is a U.S. national stage of Application No. PCT/JP2004/004554 filed

March 30, 2004.

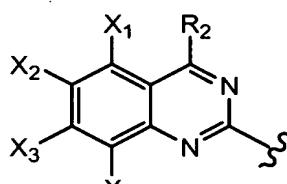
**Please replace the paragraph bridging pages 4 to 14, specifically line 18 on page 14, with the following amended paragraph:**

One aspect of the present invention relates to certain substituted heterocyclic compounds represented by Formula (I):

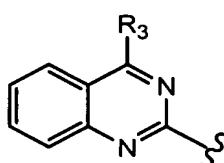


(I)

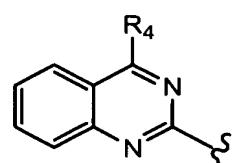
wherein Q is:



(IIa)



(IIb)



(IIc)

R<sup>1</sup> is selected from the group consisting of:

(i) C<sub>1-8</sub> alkyl, and

C<sub>1-8</sub> alkyl substituted by substituent(s) independently selected from the group consisting of:

•oxo,

•halogen,

•C<sub>1-5</sub> alkoxy carbonyl,

•C<sub>1-5</sub> alkoxy,

- C<sub>1-5</sub> alkoxy substituted by carbocyclic aryl,
- mono-C<sub>1-5</sub> alkylamino,
- mono-C<sub>1-5</sub> alkylamino substituted by carbocyclic aryl,
- di-C<sub>1-5</sub> alkylamino,
- di-C<sub>1-5</sub> alkylamino substituted by carbocyclic aryl,
- C<sub>1-5</sub> alkylthio,
- C<sub>3-6</sub> cycloalkyl,
- C<sub>3-6</sub> cycloalkyl substituted by C<sub>1-5</sub> alkyl,
- C<sub>3-6</sub> cycloalkenyl,
- carbocyclyl,
- carbocyclic aryl,
- carbocyclic aryl substituted by substituent(s) independently selected from the

group consisting of:

- hydroxy,
- halogen,
- nitro,
- amino,
- C<sub>1-5</sub> alkylcarbonylamino,
- C<sub>3-6</sub> cycloalkylcarbonylamino,
- carbocyclic aryl,
- C<sub>1-5</sub> alkyl,
- C<sub>1-5</sub> alkyl substituted by halogen,
- C<sub>1-5</sub> alkylsulfonyl,

- C<sub>2-6</sub> alkenyl,
- C<sub>1-5</sub> alkoxy, and
- C<sub>1-5</sub> alkoxy substituted by halogen,
- mono-carbocyclic arylamino,
- mono-carbocyclic arylamino substituted by substituent(s) independently selected from the group consisting of:
  - halogen,
  - C<sub>1-5</sub> alkyl,
  - C<sub>1-5</sub> alkyl substituted by halogen,
  - C<sub>1-5</sub> alkoxy, and
  - C<sub>1-5</sub> alkoxy substituted by halogen,
  - di-carbocyclic arylamino,
  - di-carbocyclic arylamino substituted by substituent(s) independently selected from the group consisting of:
    - halogen,
    - C<sub>1-5</sub> alkyl,
    - C<sub>1-5</sub> alkyl substituted by halogen,
    - C<sub>1-5</sub> alkoxy, and
    - C<sub>1-5</sub> alkoxy substituted by halogen,
    - carbocyclic aryloxy,
    - carbocyclic aryloxy substituted by substituent(s) independently selected from the group consisting of:
      - halogen,

- C<sub>1-5</sub> alkyl,
- C<sub>1-5</sub> alkyl substituted by halogen,
- C<sub>1-5</sub> alkoxy,
- C<sub>1-5</sub> alkoxy substituted by halogen, and
- carbocyclic aryl,
- hydroxy,
- heterocyclyl, and
- heterocyclyl substituted by halogen,
- (ii) C<sub>2-5</sub> alkenyl, and  
C<sub>2-5</sub> alkenyl substituted by substituent(s) independently selected from the group consisting of:
  - oxo, and
  - carbocyclic aryl,
- (iii) C<sub>2-5</sub> alkynyl,
- (iv) C<sub>3-12</sub> cycloalkyl, and  
C<sub>3-12</sub> cycloalkyl substituted by carbocyclic aryl,
- (v) carbocyclyl, and  
carbocyclyl substituted by substituent(s) independently selected from the group consisting of:
  - hydroxy, and
  - carbocyclic aryl,
- (vi) carbocyclic aryl, and

carbocyclic aryl substituted by substituent(s) independently selected from the group consisting of:

•halogen,

•cyano,

•nitro,

•amino,

•C<sub>1-10</sub> alkyl,

•C<sub>1-10</sub> alkyl substituted by substituent(s) independently selected from the group consisting of:

••halogen,

••oxo, and

••carbocyclic aryl,

•carboxy,

•C<sub>1-5</sub> alkoxy carbonyl,

•C<sub>1-7</sub> alkoxy,

•C<sub>1-7</sub> alkoxy substituted by substituent(s) independently selected from the group consisting of:

••halogen, and

••carbocyclic aryl,

•C<sub>3-6</sub> cycloalkoxy,

•carbocyclic aryloxy,

•carbocyclic aryloxy substituted by substituent(s) independently selected from the group consisting of:

- halogen,
- nitro,
- C<sub>1-5</sub> alkyl,
- C<sub>1-5</sub> alkyl substituted by halogen,
- C<sub>1-5</sub> alkoxy, and
- C<sub>1-5</sub> alkoxy substituted by halogen,
- heterocyclyloxy,
- heterocyclyloxy substituted by substituent(s) independently selected from the group consisting of:
  - halogen,
  - nitro,
  - C<sub>1-5</sub> alkyl,
  - C<sub>1-5</sub> alkyl substituted by halogen,
  - C<sub>1-5</sub> alkoxy, and
  - C<sub>1-5</sub> alkoxy substituted by halogen,
- mono-C<sub>1-5</sub> alkylamino,
- di-C<sub>1-5</sub> alkylamino,
- C<sub>1-5</sub> alkylcarbonylamino,
- C<sub>3-6</sub> cycloalkylcarbonylamino,
- C<sub>1-5</sub> alkoxy carbonylamino,
- carbocyclic aryl azo,
- carbocyclic aryl azo substituted by substituent(s) independently selected from the group consisting of:

••mono-C<sub>1-5</sub> alkylamino, and

••di-C<sub>1-5</sub> alkylamino,

•C<sub>1-5</sub> alkylthio,

•C<sub>1-5</sub> alkylthio substituted by halogen,

•carbocyclic arylthio,

•carbocyclic arylthio substituted by nitro,

•amino sulfonyl,

•heterocyclyl sulfonyl,

•C<sub>3-6</sub> cycloalkyl,

•C<sub>3-6</sub> cycloalkyl substituted by C<sub>1-5</sub> alkyl,

•carbocyclic aryl,

•carbocyclic aryl substituted by C<sub>1-5</sub> alkoxy,

•hydroxy,

•heterocyclyl, and

•heterocyclyl substituted by C<sub>1-5</sub> alkyl,

(vii) heterocyclyl, and

heterocyclyl substituted by substituent(s) independently selected from the

group consisting of:

•halogen,

•C<sub>1-5</sub> alkyl,

•C<sub>1-5</sub> alkyl substituted by halogen,

•C<sub>1-5</sub> alkoxy,

•C<sub>1-5</sub> alkoxy substituted by halogen,

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•C<sub>1-5</sub> alkoxy carbonyl,  
•C<sub>1-5</sub> alkoxy carbonyl substituted by carbocyclic aryl,  
•carbocyclic aryloxy,  
•carbocyclic aryloxy substituted by substituent(s) independently selected from  
the group consisting of:

••halogen,  
••nitro,  
••cyano,  
••hydroxy,  
••C<sub>1-5</sub> alkyl,  
••C<sub>1-5</sub> alkyl substituted by halogen,  
••mono-C<sub>1-5</sub> alkylamino,  
••di-C<sub>1-5</sub> alkylamino,  
••C<sub>1-5</sub> alkylcarbonylamino,  
••C<sub>3-6</sub> cycloalkylcarbonylamino,  
••C<sub>1-5</sub> alkoxy,  
••C<sub>1-5</sub> alkoxy substituted by halogen,  
••C<sub>3-6</sub> cycloalkyl,  
••C<sub>2-5</sub> alkenyl,  
••C<sub>2-5</sub> alkynyl,  
••carboxy,  
••C<sub>1-5</sub> alkoxy carbonyl,  
••mono-C<sub>1-5</sub> alkylaminocarbonyl,

- di-C<sub>1-5</sub> alkylaminocarbonyl,
- mono-C<sub>3-6</sub> cycloalkylaminocarbonyl,
- di-C<sub>3-6</sub> cycloalkylaminocarbonyl,
- mono-C<sub>1-5</sub> alkylaminocarbonylamino,
- di-C<sub>1-5</sub> alkylaminocarbonylamino,
- mono-C<sub>3-6</sub> cycloalkylaminocarbonylamino,
- di-C<sub>3-6</sub> cycloalkylaminocarbonylamino,
- C<sub>1-5</sub> alkylthio,
- C<sub>1-5</sub> alkylthio substituted by halogen,
- C<sub>1-5</sub> alkylsulfinyl,
- C<sub>1-5</sub> alkylsulfinyl substituted by halogen,
- C<sub>1-5</sub> alkylsulfonyl, and
- C<sub>1-5</sub> alkylsulfonyl substituted by halogen,
- heterocyclyloxy,
- heterocyclyloxy substituted by substituent(s) independently selected from the group consisting of:

- halogen,
- nitro,
- C<sub>1-5</sub> alkyl,
- C<sub>1-5</sub> alkyl substituted by halogen,
- C<sub>1-5</sub> alkoxy, and
- C<sub>1-5</sub> alkoxy substituted by halogen,
- carbocyclic aryl, and

•heterocyclyl;

R<sub>2</sub> is C<sub>1-5</sub> alkyl or -N(R<sub>2a</sub>)(R<sub>2b</sub>); wherein R<sub>2a</sub> and R<sub>2b</sub> are independently hydrogen or C<sub>1-5</sub> alkyl;

R<sub>3</sub> is C<sub>1-5</sub> alkyl;

R<sub>4</sub> is -NHNH<sub>2</sub>, -NHNHBoc, -N(R<sub>4a</sub>)(R<sub>4b</sub>), morpholino, 4-acetyl-piperazyl, or 4-phenyl-piperazyl; wherein R<sub>4a</sub> is hydrogen or C<sub>1-5</sub> alkyl; R<sub>4b</sub> is C<sub>1-5</sub> alkyl, C<sub>1-5</sub> alkyl substituted by substituent(s) independently selected from the group consisting of:

•hydroxy,

•C<sub>1-5</sub> alkoxy,

•amino,

•-NHBoc,

•C<sub>3-6</sub> cycloalkyl,

•carbocyclic aryl,

•carbocyclic aryl substituted by substituent(s) independently selected from the group consisting of:

•halogen,

•C<sub>1-5</sub> alkyl,

•C<sub>1-5</sub> alkoxy, and

•-SO<sub>2</sub>NH<sub>2</sub>, and

•heterocyclyl,

C<sub>3-6</sub> cycloalkyl, carbocyclic aryl, carbocyclic aryl substituted by substituent(s) independently selected from the group consisting of:

•halogen,

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•C<sub>1-5</sub> alkyl,

•C<sub>1-5</sub> alkoxy, and

a group of Formula (III):



(III) ;

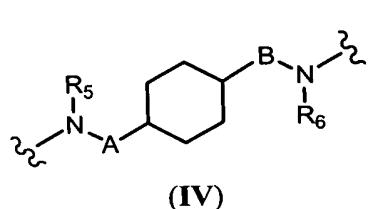
wherein Boc is carbamic acid *tert*-butyl ester and G is C<sub>1-5</sub> alkyl or C<sub>1-5</sub> alkoxy substituted by substituent(s) independently selected from the group consisting of:

•carbocyclic aryl,

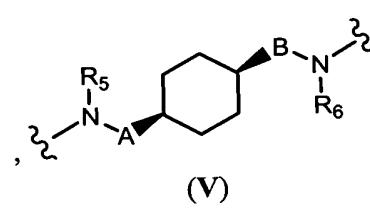
•halogenated carbocyclic aryl, and

•carbocyclic aryl substituted by C<sub>1-5</sub> alkoxy;

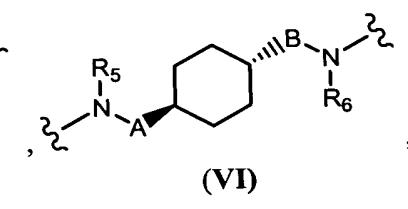
L is selected from the group consisting of Formulae (IV) to (XIX):



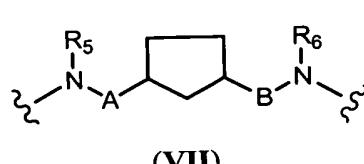
(IV)



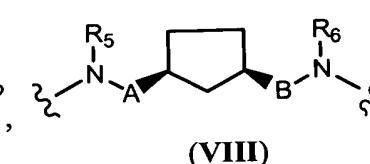
(V)



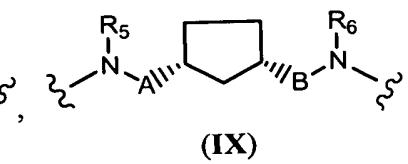
(VI)



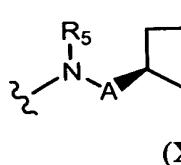
(VII)



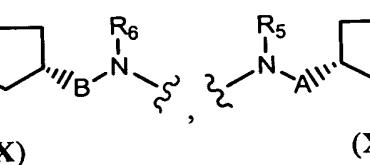
(VIII)



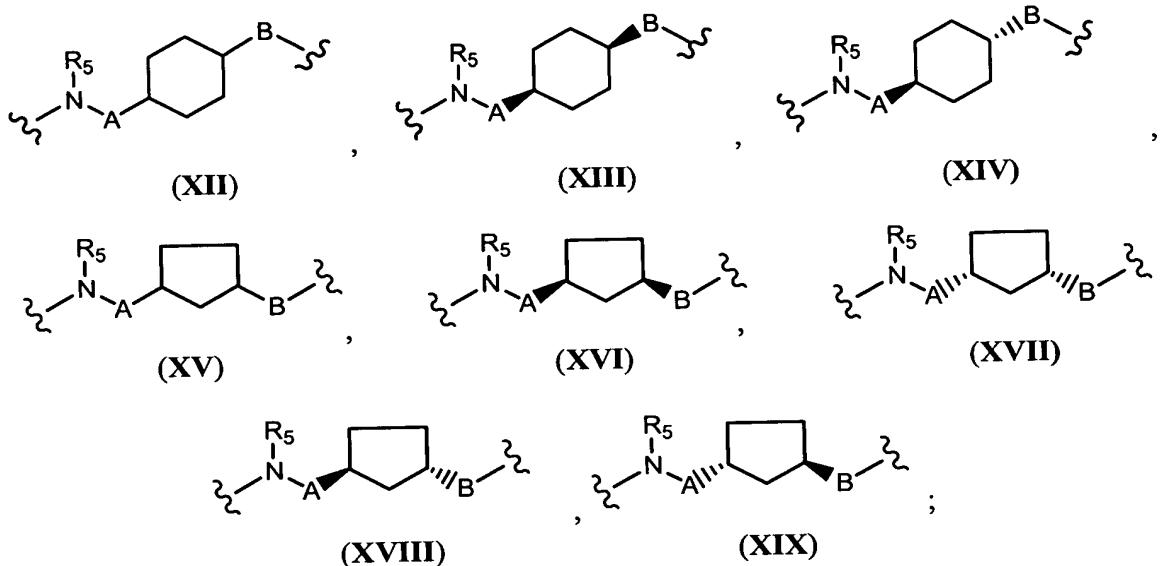
(IX)



(X)



(XI)



wherein  $R_5$  and  $R_6$  are independently hydrogen or  $C_{1-5}$  alkyl; and  $A$  and  $B$  are independently a single bond,  $-CH_2-$ , or  $-(CH_2)_2-$ ;  
 $X_1$ ,  $X_2$ ,  $X_3$  and  $X_4$  are independently selected from the group consisting of hydrogen, halogen,  $C_{1-4}$  alkyl,  $C_{1-4}$  alkyl substituted by halogen,  $C_{1-4}$  alkylthio,  $C_{1-4}$  alkylsulfinyl,  $C_{1-4}$  alkylsulfonyl,  $C_{1-4}$  alkoxy,  $C_{1-4}$  alkoxy substituted by halogen, nitro, amino, mono- $C_{1-4}$  alkylamino, di- $C_{1-4}$  alkylamino, piperidyl, morpholinyl, mono- $C_{1-4}$  alkylaminosulfonyl, di- $C_{1-4}$  alkylaminosulfonyl and hydroxy; provided that at least one substituent selected from the group consisting of  $X_1$ ,  $X_2$ ,  $X_3$  and  $X_4$  is not hydrogen; and

$Y$  is selected from the group consisting of:

- (i)  $-C(O)NR_7-$ ,  $-C(S)NR_7-$ , or  $-C(O)O-$  when  $L$  is selected from the group consisting of Formulae (IV) to (XIX); wherein  $R_7$  is hydrogen or  $C_{1-5}$  alkyl;
- (ii)  $-S(O)_2-$ ,  $-C(O)-$ , a single bond or  $-CH_2-$  when  $L$  is selected from the group consisting of Formulae (IV) to (XI), and  $Q$  is Formula (IIa) or (IIb);

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(iii) -S(O)<sub>2</sub>-, -C(O)-, a single bond or -CH<sub>2</sub>- when L is selected from the group consisting of Formulae (VII) to (XI), and Q is Formula (IIc); and

(iv) -OC(O)- when L is selected from the group consisting of Formulae (XII) to (XIX);

wherein carbocyclic aryl is phenyl, naphthyl, or biphenyl;

carbocyclyl is indanyl, bicyclo[2.2.1]heptyl, bicyclo[2.2.1]heptenyl, adamantyl, adamantyl, 9H-fluorenyl, menthyl, 1,2,3,4-tetrahydro-naphthalen-1-yl, or 1H-indolyl; heterocyclyl is 2,3-dihydro-benzo[1,4]dioxinyl, 3,4-dihydro-2H-benzo[b][1,4]dioxepinyl, 4,5,6,7-tetrahydro-benzo[b]thienyl, 4H-benzo[1,3]dioxinyl, benzo[1,3]dioxolyl, benzo[2,1,3]thiadiazolyl, benzothiazolyl, furyl, isoxazolyl, morpholinyl, oxazolyl, piperidyl, pyrazolyl, pyridyl, tetrahydrofuryl, thienyl, dibenzofuranyl, 1H-benzoimidazolyl, or thiazolyl; and

halogen is fluoro, chloro, bromo, or iodo;

or a pharmaceutically acceptable salt, hydrate or solvate thereof.

**Please replace the paragraph bridging pages 17-21, specifically line 19 on page 21, with the following amended paragraph:**

In some embodiments of the present invention, Q is Formulae (IIa), (IIb), or (IIc);

R<sub>1</sub> is selected from the group consisting of:

(i) C<sub>1-8</sub> alkyl, and

C<sub>1-8</sub> alkyl substituted by substituent(s) independently selected from the group consisting of:

•halogen,

•C<sub>1-5</sub> alkoxy carbonyl,

•C<sub>1-5</sub> alkoxy,

•C<sub>1-5</sub> alkoxy substituted by carbocyclic aryl,

•mono-C<sub>1-5</sub> alkylamino,

•di-C<sub>1-5</sub> alkylamino,

•C<sub>3-6</sub> cycloalkyl,

•C<sub>3-6</sub> cycloalkenyl,

•carbocyclyl,

•carbocyclic aryl,

•carbocyclic aryl substituted by substituent(s) independently selected from the group consisting of:

••hydroxy,

••halogen,

••nitro,

••C<sub>1-5</sub> alkylcarbonylamino,

••C<sub>3-6</sub> cycloalkylcarbonylamino,  
••C<sub>1-5</sub> alkyl,  
••C<sub>1-5</sub> alkyl substituted by halogen,  
••C<sub>1-5</sub> alkylsulfonyl,  
••C<sub>2-6</sub> alkenyl,  
••C<sub>1-5</sub> alkoxy,  
••C<sub>1-5</sub> alkoxy substituted by halogen, and  
••carbocyclic aryl,

•heterocyclyl, and

•heterocyclyl substituted by halogen,

(ii) C<sub>2-5</sub> alkenyl, and

C<sub>2-5</sub> alkenyl substituted by carbocyclic aryl,

(iii) C<sub>2-5</sub> alkynyl,

(iv) C<sub>3-12</sub> cycloalkyl, and

C<sub>3-12</sub> cycloalkyl substituted by carbocyclic aryl,

(v) carbocyclyl, and

carbocyclyl by substituent(s) independently selected from the group consisting

of:

•hydroxy, and

•carbocyclic aryl,

(vi) carbocyclic aryl, and

carbocyclic aryl substituted by substituent(s) independently selected from the

group consisting of:

- halogen,
- cyano,
- nitro,
- C<sub>1-10</sub> alkyl,
- C<sub>1-10</sub> alkyl substituted by substituent(s) independently selected from the group consisting of:

- carboxy,
- C<sub>1-5</sub> alkoxy carbonyl,
- C<sub>1-7</sub> alkoxy,
- C<sub>1-7</sub> alkoxy substituted by substituent(s) independently selected from the group consisting of:

- halogen, and
- carbocyclic aryl,
- carbocyclic aryloxy,
- carbocyclic aryloxy substituted by nitro,
- mono-C<sub>1-5</sub> alkylamino,
- di-C<sub>1-5</sub> alkylamino,
- C<sub>1-5</sub> alkoxy carbonylamino,
- carbocyclic aryl azo,

• carbocyclic aryl azo substituted by substituent(s) independently selected from  
the group consisting of:

- mono-C<sub>1-5</sub> alkylamino, and
- di-C<sub>1-5</sub> alkylamino,
- C<sub>1-5</sub> alkylthio,
- C<sub>1-5</sub> alkylthio substituted by halogen,
- carbocyclic arylthio,
- carbocyclic arylthio substituted by nitro,
- amino sulfonyl,
- heterocyclyl sulfonyl,
- C<sub>3-6</sub> cycloalkyl,
- C<sub>3-6</sub> cycloalkyl substituted by C<sub>1-5</sub> alkyl,
- carbocyclic aryl,
- heterocyclyl, and
- heterocyclyl substituted by C<sub>1-5</sub> alkyl,

(vii) heterocyclyl, and

heterocyclyl substituted by substituent(s) independently selected from the  
group consisting of:

- halogen,
- C<sub>1-5</sub> alkyl,
- C<sub>1-5</sub> alkyl substituted by halogen,
- C<sub>1-5</sub> alkoxy,
- C<sub>1-5</sub> alkoxy carbonyl,

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- C<sub>1-5</sub> alkoxy carbonyl substituted by carbocyclic aryl,
- carbocyclic aryloxy,
- carbocyclic aryl, and
- heterocyclyl;

R<sub>2</sub> is -N(R<sub>2a</sub>)(R<sub>2b</sub>), wherein R<sub>2a</sub> is hydrogen or C<sub>1-5</sub> alkyl; R<sub>2b</sub> is C<sub>1-5</sub> alkyl;

R<sub>3</sub> is C<sub>1-5</sub> alkyl;

R<sub>4</sub> is -N(R<sub>4a</sub>)(R<sub>4b</sub>) wherein R<sub>4a</sub> is hydrogen or C<sub>1-5</sub> alkyl; R<sub>4b</sub> is C<sub>1-5</sub> alkyl;

L is selected from Formula (V), (VIII), (IX), (XIII), (XVI), or (XVII);

X<sub>1</sub>, X<sub>2</sub>, X<sub>3</sub> and X<sub>4</sub> are independently selected from the group consisting of hydrogen, halogen, and C<sub>1-4</sub> alkyl; provided that at least one substituent selected from the group consisting of X<sub>1</sub>, X<sub>2</sub>, X<sub>3</sub> and X<sub>4</sub> is not hydrogen; and

Y is selected from the group consisting of:

(i) -C(O)NR<sub>7-</sub>, -C(S)NR<sub>7-</sub>, or -C(O)O- when L is selected from the group consisting of Formula (V), (VIII), (IX), (XIII), (XVI), or (XVII); wherein R<sub>7</sub> is hydrogen or C<sub>1-5</sub> alkyl;

(ii) -S(O)<sub>2</sub>-, -C(O)-, a single bond or -CH<sub>2</sub>- when L is selected from the group consisting of Formula (VIII) or (IX); and

(iii) -OC(O)- when L is selected from the group consisting of Formula (XIII), (XVI), or (XVII);

wherein carbocyclic aryl is phenyl or naphthyl;  
 carbocyclyl is indanyl, bicyclo[2.2.1]heptyl, bicyclo[2.2.1]heptenyl,  
adamantyl, adamantyl, 9H-fluorenyl, menthyl, 1,2,3,4-tetrahydro-naphthalen-1-yl, or 1*H*-indolyl;

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heterocyclyl is 2,3-dihydro-benzo[1,4]dioxinyl, 3,4-dihydro-2*H*-  
benzo[b][1,4]dioxepinyl, 4,5,6,7-tetrahydro-benzo[b]thienyl, 4*H*-benzo[1,3]dioxinyl,  
benzo[1,3]dioxolyl, benzo[2,1,3]thiadiazolyl, benzothiazolyl, furyl, isoxazolyl, morpholinyl,  
oxazolyl, piperidyl, pyrazolyl, pyridyl, tetrahydrofuryl, thienyl, dibenzofuranyl, 1*H*-  
benzoimidazolyl, or thiazolyl; and  
halogen is fluoro, chloro, bromo, or iodo;  
or a pharmaceutically acceptable salt, hydrate or solvate thereof.

**Please replace the paragraph bridging pages 22 to 24, specifically line 10 on page 24, with the following amended paragraph:**

In some embodiments of the present invention, R<sub>1</sub> is selected from the group consisting of:

(i) C<sub>1-5</sub> alkyl, and

C<sub>1-5</sub> alkyl substituted by substituent(s) independently selected from the group

consisting of:

•C<sub>1-5</sub> alkoxy carbonyl,

•carbocyclic aryl,

•carbocyclic aryl substituted by substituent(s) independently selected from the

group consisting of:

••halogen,

••C<sub>1-5</sub> alkyl,

••C<sub>2-5</sub> alkenyl, and

••C<sub>1-5</sub> alkoxy,

•C<sub>1-5</sub> alkylthio, and

•heterocyclyl,

(ii) C<sub>3-6</sub> cycloalkyl, and

C<sub>3-6</sub> cycloalkyl substituted by carbocyclic aryl,

(iii) carbocyclyl,

(iv) carbocyclic aryl, and

carbocyclic aryl substituted by substituent(s) independently selected from the

group consisting of:

- halogen,
- cyano,
- nitro,
- C<sub>1-5</sub> alkyl,
- C<sub>1-5</sub> alkyl substituted by substituent(s) independently selected from the group consisting of:

- halogen,
- oxo, and
- carbocyclic aryl,
- C<sub>1-5</sub> alkoxy carbonyl,
- C<sub>1-7</sub> alkoxy,
- C<sub>1-7</sub> alkoxy substituted by substituent(s) independently selected from the group consisting of:

- halogen, and
- carbocyclic aryl,
- cycloalkoxy,
- carbocyclic aryloxy,
- mono-C<sub>1-5</sub> alkylamino,
- di-C<sub>1-5</sub> alkylamino,
- C<sub>1-5</sub> alkylthio,
- C<sub>1-5</sub> alkylthio substituted by halogen,
- carbocyclic aryl,
- heterocyclyl, and

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•heterocyclyl substituted by C<sub>1-5</sub> alkyl,  
(v) heterocyclyl, and  
heterocyclyl substituted by substituent(s) independently selected from the  
group consisting of:

- halogen,
- C<sub>1-5</sub> alkyl,
- C<sub>1-5</sub> alkyl substituted by halogen,
- C<sub>1-5</sub> alkoxy carbonyl
- C<sub>1-5</sub> alkoxy carbonyl substituted by carbocyclic aryl, and
- carbocyclic aryl;

L is Formula (V);

and

Y is -C(O)NR<sub>7</sub>-; wherein R<sub>7</sub> is hydrogen or C<sub>1-5</sub> alkyl;  
wherein carbocyclic aryl is phenyl or naphthyl;  
carbocyclyl is indanyl, ~~adamantly~~adamantyl, or 9*H*-fluorenyl;  
heterocyclyl is 2,3-dihydro-benzo[1,4]dioxinyl, 3,4-dihydro-2*H*-  
benzo[b][1,4]dioxepinyl, 4*H*-benzo[1,3]dioxinyl, benzo[1,3]dioxolyl, benzothiazolyl,  
furyl, isoxazolyl, piperidyl, pyridyl, or thienyl; and  
halogen is fluoro, chloro, bromo, or iodo;  
or a pharmaceutically acceptable salt, hydrate or solvate thereof.

**Please replace the paragraph bridging pages 50 to 53, specifically line 10 on page 53, with the following amended paragraph:**

In some of the embodiments of the present invention, R<sub>1</sub> is selected from the group consisting of:

(i) C<sub>1-8</sub> alkyl, and

C<sub>1-8</sub> alkyl substituted by substituent(s) independently selected from the group

consisting of:

•mono-C<sub>1-5</sub> alkylamino,

•di-C<sub>1-5</sub> alkylamino,

•C<sub>3-6</sub> cycloalkyl,

•C<sub>3-6</sub> cycloalkenyl,

•carbocyclic aryl,

•carbocyclic aryl substituted by substituent(s) independently selected from the

group consisting of:

•halogen,

•C<sub>1-5</sub> alkyl, and

•C<sub>1-5</sub> alkoxy,

•heterocyclyl,

(ii) C<sub>2-5</sub> alkynyl,

(iii) C<sub>2-5</sub> alkenyl, and

C<sub>2-5</sub> alkenyl substituted by carbocyclic aryl,

(iv) C<sub>3-12</sub> cycloalkyl,

(v) carbocyclyl,

(vi) carbocyclic aryl, and

carbocyclic aryl substituted by substituent(s) independently selected from the group consisting of:

•halogen,

•cyano,

•nitro,

•C<sub>1-10</sub> alkyl,

•C<sub>1-10</sub> alkyl substituted by substituent(s) independently selected from the group

consisting of:

••halogen, and

••OXO,

•carboxy,

•C<sub>1-5</sub> alkoxy carbonyl,

•C<sub>1-5</sub> alkoxy,

•C<sub>1-5</sub> alkoxy substituted by substituent(s) independently selected from the

group consisting of:

••halogen, and

••carbocyclic aryl,

•carbocyclic aryloxy,

•carbocyclic aryloxy substituted by nitro,

•mono-C<sub>1-5</sub> alkylamino,

•di-C<sub>1-5</sub> alkylamino,

•C<sub>1-5</sub> alkoxy carbonylamino,

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• carbocyclic aryl azo,  
• carbocyclic aryl azo substituted by substituent(s) independently selected from  
the group consisting of:

- mono-C<sub>1-5</sub> alkylamino, and
- di-C<sub>1-5</sub> alkylamino,
- C<sub>1-5</sub> alkylthio,
- C<sub>1-5</sub> alkylthio substituted by halogen,
- carbocyclic arylthio,
- carbocyclic arylthio substituted by nitro,
- amino sulfonyl,
- heterocyclyl sulfonyl,
- C<sub>3-6</sub> cycloalkyl,
- C<sub>3-6</sub> cycloalkyl substituted by C<sub>1-5</sub> alkyl,
- carbocyclic aryl, and
- heterocyclyl,

(vii) heterocyclyl, and  
heterocyclyl substituted by substituent(s) independently selected from the

group consisting of:

- C<sub>1-5</sub> alkyl,
- C<sub>1-5</sub> alkoxy carbonyl,
- carbocyclic aryloxy,
- carbocyclic aryl, and
- heterocyclyl;

L is Formula (V); and

Y is -C(S)NR<sub>7</sub>-; wherein R<sub>7</sub> is hydrogen or C<sub>1-5</sub> alkyl;

wherein carbocyclic aryl is phenyl or naphthyl;

carbocyclyl is indanyl, bicyclo[2.2.1]heptyl, bicyclo[2.2.1]heptenyl, or

adamantlyadamantyl;

heterocyclyl is 2,3-dihydro-benzo[1,4]dioxinyl, 4,5,6,7-tetrahydro-

benzo[b]thienyl, benzo[1,3]dioxolyl, benzo[2,1,3]thiadiazolyl, furyl, isoxazolyl,

morpholinyl, oxazolyl, piperidyl, pyrazolyl, pyridyl, tetrahydrofuryl, or thienyl; and

halogen is fluoro, chloro, bromo, or iodo;

or a pharmaceutically acceptable salt, hydrate or solvate thereof.

**Please replace the last paragraph on page 266 with the following amended**

**paragraph:**

293 cells (human kidney, ATCC), transiently transfected with 10  $\mu$ g human MCH receptor and 60  $\mu$ l Lipofectamine (per 15-cm dish), are grown in the dish for 24 hours (75% confluence) with a media change and removed with 10 ml/dish of Hepes-EDTA buffer (20mM Hepes + 10 mM EDTA, pH 7.4). The cells are then centrifuged in a Beckman Coulter centrifuge for 20 minutes, 17,000 rpm (JA-25.50 rotor). Subsequently, the pellet is resuspended in 20 mM Hepes + 1 mM EDTA, pH 7.4 and homogenized with a 50- ml Dounce homogenizer and again centrifuged. After removing the supernatant, the pellets can be stored at -80°C, until used in binding assay. When used in the assay, membranes are thawed on ice for 20 minutes and then 10 mL of incubation buffer (20 mM Hepes, 1 mM MgCl<sub>2</sub>, 100 mM NaCl, pH 7.4) added. The membranes are then vortexed to resuspend the crude membrane pellet and homogenized with a Brinkmann PT-3100 Polytron homogenizer for 15 seconds at setting 6. The concentration of membrane protein is determined using the BRL Bradford protein assay.

**Please replace the first paragraph on page 267 with the following amended paragraph:**

For total binding, a total volume of 50 $\mu$ l of appropriately diluted membranes (diluted in assay buffer containing 50mM Tris HCl (pH 7.4), 10mM MgCl<sub>2</sub>, and 1mM EDTA; 5-50 $\mu$ g $\mu$ g protein) is added to 96-well polypropylene microtiter plates followed by addition of 100 $\mu$ l of assay buffer and 50 $\mu$ l of **Radiolabelled MCH Ligand**. For nonspecific binding, 50 $\mu$ l of assay buffer is added instead of 100 $\mu$ l and an additional 50 $\mu$ l of 10 $\mu$ M $\mu$ M cold MCH is added before 50 $\mu$ l of **Radiolabelled MCH Ligand** is added. Plates are then incubated at room temperature for 60-120 minutes. The binding reaction is terminated by filtering assay plates through a Microplate Devices GF/C Unifilter filtration plate with a Brandell 96-well plate harvester followed by washing with cold 50 mM Tris HCl, pH 7.4 containing 0.9% NaCl. Then, the bottom of the filtration plate are sealed, 50  $\mu$ l of Optiphase Supermix is added to each well, the top of the plates are sealed, and plates are counted in a Trilux MicroBeta scintillation counter. For compound competition studies, instead of adding 100  $\mu$ l of assay buffer, 100  $\mu$ l of appropriately diluted test compound is added to appropriate wells followed by addition of 50  $\mu$ l of Radiolabelled MCH Ligand.